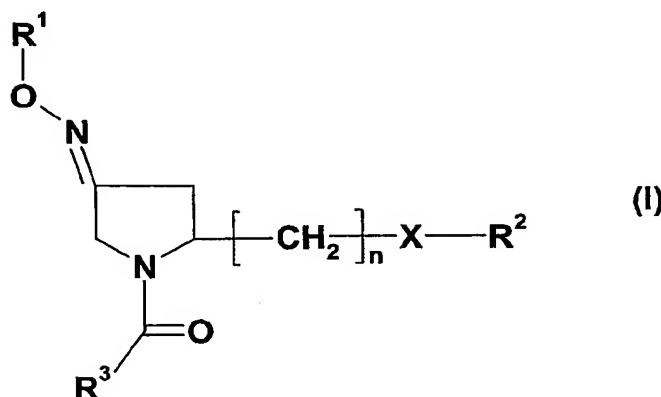


Claims

1. A pyrrolidine derivative of Formula I:



its geometrical isomers, its optically active forms as enantiomers, diastereomers, mixtures of these and its racemate forms, as well as salts thereof, wherein:

R¹ is selected from the group comprising or consisting of H and C₁-C₆-alkyl;

R² is selected from the group comprising or consisting of hydrogen, C₁-C₆-alkyl, C₁-C₆-alkyl aryl, heteroaryl, C₁-C₆-alkyl heteroaryl, C₂-C₆-alkenyl, C₂-C₆-alkenyl aryl, C₂-C₆-alkenyl heteroaryl, C₂-C₆-alkynyl, C₂-C₆-alkynyl aryl, C₂-C₆-alkynyl heteroaryl, C₃-C₈-cycloalkyl, heterocycloalkyl, C₁-C₆-alkyl cycloalkyl, C₁-C₆-alkyl heterocycloalkyl, C₁-C₆-alkyl carboxy, acyl, C₁-C₆-alkyl acyl, C₁-C₆-alkyl acyloxy, C₁-C₆-alkyl alkoxy, alkoxy carbonyl, C₁-C₆-alkyl alkoxy carbonyl, aminocarbonyl, C₁-C₆-alkyl aminocarbonyl, C₁-C₆-alkyl acylamino, C₁-C₆-alkyl ureido, amino, C₁-C₆-alkyl amino, sulfonyloxy, C₁-C₆-alkyl sulfonyloxy, sulfonyl, C₁-C₆-alkyl sulfonyl, sulfinyl, C₁-C₆-alkyl sulfinyl, C₁-C₆-alkyl sulfanyl, C₁-C₆-alkyl sulfonylamino;

R³ is selected from the group comprising or consisting of aryl and heteroaryl;

X is selected from the group consisting of O or NR⁴;

R⁴ is selected from the group comprising or consisting of H, C₁-C₆-alkyl, C₁-C₆-alkyl aryl, C₁-C₆-alkyl heteroaryl, aryl, heteroaryl; or

R² and R⁴ can form together with the N atom to which they are linked to, a 5-8 membered saturated or unsaturated heterocycloalkyl ring; and

n is an integer from 1 to 3.

2. A pyrrolidine derivative according to claim 1, wherein R¹ is methyl.

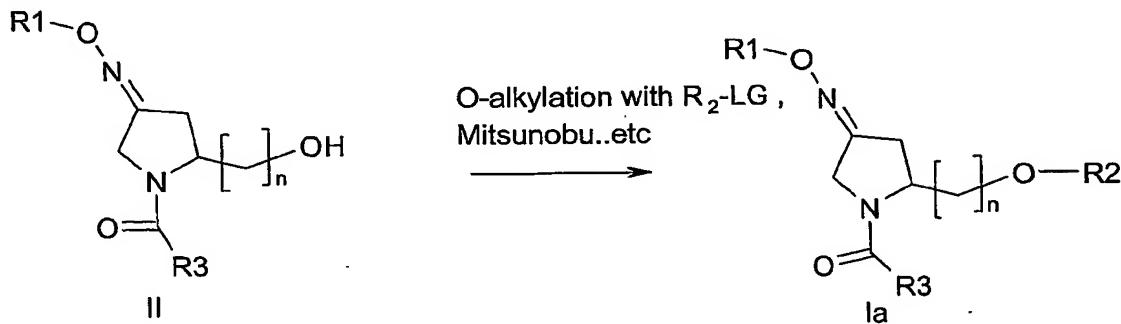
3. A pyrrolidine derivative according to claim 1 or 2, wherein R³ is a phenyl.

4. A pyrrolidine derivative according to any of the preceding claims, wherein n is an integer 1 or 2.
5. A pyrrolidine derivative according to any of the preceding claims wherein R² and R⁴ form together with the N atom to which they are linked, a 5 or 6 membered cycloalkyl or heterocycloalkyl ring;
6. A pyrrolidine derivative according to claims 1 to 4 wherein X is O or NH.
- 10 7. A pyrrolidine derivative according to any of the preceding claims selected from the following group:
(3EZ,5S)-5-(hydroxymethyl)-1-[(2'-methyl-1,1'-biphenyl-4-yl)carbonyl]pyrrolidin-3-one O-methyloxime;
(3EZ,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-(hydroxymethyl)pyrrolidin-3-one O-methyloxime;
15 (3E,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-(hydroxymethyl)pyrrolidin-3-one O-methyloxime;
(3Z,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-[(4-methylpiperazin-1-yl)methyl]pyrrolidin-3-one O-methyloxime;
20 tert-butyl {[(2S,4EZ)-1-(1,1'-biphenyl-4-ylcarbonyl)-4-(methoxyimino)pyrrolidin-2-yl]methoxy} acetate;
{ [(2S,4EZ)-1-(1,1'-biphenyl-4-ylcarbonyl)-4-(methoxyimino)pyrrolidin-2-yl]-methoxy} acetic acid;
25 2- {[(2S,4EZ)-1-(1,1'-biphenyl-4-ylcarbonyl)-4-(methoxyimino)pyrrolidin-2-yl]-methoxy}-N-(2-pyrrolidin-1-ylethyl)acetamide;
(3EZ,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-(methoxymethyl)pyrrolidin-3-one O-methyloxime;
30 (3EZ,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-[(4-methylpiperazin-1-yl)methyl]-pyrrolidin-3-one O-methyloxime;
(3EZ,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-{[(4-methoxyphenyl)amino]methyl}-pyrrolidin-3-one O-methyloxime;
(3EZ,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-({[2-(1H-pyrazol-1-yl)ethyl]amino}-methyl)-pyrrolidin-3-one O-methyloxime;

2-{[(2S,4EZ)-1-(1,1'-biphenyl-4-ylcarbonyl)-4-(methoxyimino)pyrrolidin-2-yl]-methyl}-1H-isoindole-1,3(2H)-dione;
(3EZ,5S)-5-(aminomethyl)-1-(1,1'-biphenyl-4-ylcarbonyl)pyrrolidin-3-one O-methyl-oxime;
5 N-{[(2S,4EZ)-1-(1,1'-biphenyl-4-ylcarbonyl)-4-(methoxyimino)pyrrolidin-2-yl]methyl}acetamide;
(3EZ,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-(piperidin-1-ylmethyl)pyrrolidin-3-one O-methyloxime;
(3EZ,5S)-1-(1,1'-biphenyl-4-ylcarbonyl)-5-(2-hydroxyethyl)pyrrolidin-3-one O-methyloxime.
10

8. A pyrrolidine according to any of the preceding claims for use as a medicament.
9. Use of a pyrrolidine derivative according to any of claims 1 to 7 as well as isomers, optically active forms as enantiomers, diastereomers and mixtures of these, as well as salts thereof for the preparation of a medicament for the prevention and/or treatment of preterm labor, premature birth or dysmenorrhea.
15
10. Use of a pyrrolidine according to claim 1 to 7, for the preparation of a medicament for the treatment of disorders requiring the modulation of the oxytocin receptor.
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11. Use according to claim 10, for the treatment or prevention of disorders associated with the oxytocin receptor activity.
- 25 12. Use according to claim 10 or 11, wherein said modulation consists in the blocking of the oxytocin receptor or in antagonising the binding of oxytocin to its receptor.
13. A pharmaceutical composition containing a pyrrolidine derivative according to any of claims 1 to 7 and a pharmaceutically acceptable carrier, diluent or excipient thereof.
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14. A process for the preparation of a pyrrolidine derivative according to any of claims 1 to 7, wherein X is O, comprising the step of an O-alkylation of alcohol derivatives

of formula (II) with an alkylating agent $R^2\text{-LG}$ wherein LG is a leaving group, with R^1 , R^2 , R^3 and n being as defined above.



15. A process for the preparation of a pyrrolidine derivative according to any of claims 1 to 7 wherein X is NR^4 , comprising the step of a reductively aminating an aldehyde derivative of formula (XI) with an amine HNR^2R^4 wherein R^1 , R^2 , R^3 , R^4 and n are defined above.

